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OM protein - protein search, using sw model

Run on: June 25, 2003, 14:20:41 ; Search time 33 Seconds

(without alignments)  
444.169 Million cell updates/sec

Title: US-09-622-613b-15

Perfect score: 602

Sequence: 1 QMNAFPOKHIIITPILICNT.....ICVKCENQYVPHAGIGRCP 110

Scoring table: BLOSUM62

Gapop 10.0, Gapext 0.5

Searched: 908470 seqs, 133250620 residues

Total number of hits satisfying chosen parameters: 908470

Minimum DB seq length: 10

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database :

1: /SID2/gcgdata/geneseq/emb1/AA1980.DAT:\*  
2: /SID2/gcgdata/geneseq/emb1/AA1981.DAT:\*  
3: /SID2/gcgdata/geneseq/emb1/AA1982.DAT:\*  
4: /SID2/gcgdata/geneseq/emb1/AA1983.DAT:\*  
5: /SID2/gcgdata/geneseq/emb1/AA1984.DAT:\*  
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9: /SID2/gcgdata/geneseq/emb1/AA1988.DAT:\*  
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19: /SID2/gcgdata/geneseq/emb1/AA1998.DAT:\*  
20: /SID2/gcgdata/geneseq/emb1/AA1999.DAT:\*  
21: /SID2/gcgdata/geneseq/emb1/AA2000.DAT:\*  
22: /SID2/gcgdata/geneseq/emb1/AA2001.DAT:\*  
23: /SID2/gcgdata/geneseq/emb1/AA2002.DAT:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length DB	ID	Description
1	602	100.0	110	AAV28872	Rana catesbeiana o
2	602	100.0	110	AAV28873	Recombinant Met(-1
3	597	99.2	110	AAV28877	Recombinant RacOR1
4	597	99.2	110	AAV28878	Recombinant Met(-1
5	596	99.0	110	AAV28874	Recombinant RacOR1
6	596	99.0	110	AAV28876	Recombinant Met(-1
7	588.5	97.8	111	AAV33321	Frog lectin protei
8	284.5	47.3	104	AAW06544	Antitumor protein
9	281.5	46.8	104	AAV28865	Rana pipiens liver
10	281.5	46.8	105	AAV28867	Recombinant Met(-1

11	281.5	46.8	127	20	AAV28879	Rana pipiens Clone
12	278.5	46.3	104	20	AAV28866	Recombinant RAPR1
13	278.5	46.3	105	20	AAV28869	Recombinant Met(-1
14	277.5	46.1	104	18	AAW30301	Recombinant onc pr
15	277.5	46.1	104	22	AAW31666	Amino acid sequenc
16	277.5	46.1	105	20	AAV39400	Recombinant frog O
17	277.5	46.1	105	18	AAW35126	R. pipiens recombi
18	276.5	45.9	104	20	AAV28870	Recombinant RAPR1
19	276.5	45.9	105	20	AAV28871	Recombinant Met(-1
20	274.5	45.6	104	12	AAW12344	Protein with activ
21	274.5	45.6	104	15	AAW47303	ONCONASE (pharmac
22	274.5	45.6	104	17	AAW0736	Protein derived fr
23	274.5	45.6	104	18	AAW06543	Antitumor protein
24	274.5	45.6	104	18	AAW14065	Onconase (RTM) pro
25	274.5	45.6	104	20	AAV33322	Frog onconase prot
26	274.5	45.6	104	20	AAW88233	Rana pipiens RNase
27	274.5	45.6	105	18	AAW35123	R. pipiens recombi
28	274.5	45.6	105	18	AAW35125	R. pipiens recombi
29	274.5	45.6	358	18	AAW35130	R. pipiens recombi
30	272.5	45.3	104	22	AAW31667	Amino acid sequenc
31	272.5	45.3	106	18	AAW35132	R. pipiens recombi
32	272.5	45.3	107	18	AAW35117	R. pipiens recombi
33	272.5	45.3	112	18	AAW35118	R. pipiens recombi
34	272.5	45.3	251	18	AAW35134	R. pipiens recombi
35	272.5	45.3	254	18	AAW35135	R. pipiens recombi
36	272.5	45.3	355	18	AAW35129	R. pipiens recombi
37	272.5	45.3	355	18	AAW35133	R. pipiens recombi
38	272.5	45.3	366	18	AAW35132	R. pipiens recombi
39	271.5	45.1	104	18	AAW30302	Recombinant onc pr
40	267.5	44.4	104	18	AAW18224	Antitumor generi
41	267.5	44.4	105	18	AAW35115	R. pipiens recombi
42	267.5	44.4	105	18	AAW35116	R. pipiens recombi
43	263.5	43.8	358	18	AAW35127	R. pipiens recombi
44	263.5	43.8	365	18	AAW35131	R. pipiens recombi
45	249.5	41.4	107	18	AAW35120	R. pipiens recombi

#### ALIGNMENTS

RESULT 1	
AAV28872	
ID	AAV28872 standard; Protein; 110 AA.
XX	
AC	AAV28872;
XX	
DT	25-JAN-2000 (first entry)
XX	
DE	Rana catesbeiana oocyte ribonuclease (RacOR1) amino acid sequence.
XX	
KW	Rana catesbeiana oocyte ribonuclease; RacOR1; covalently bound; CD22;
KW	LL2 antibody; ligand binding moiety; cancerous B cell; Kaposi's Sarcoma;
KW	human chorionic gonadotropin; hCG; recombinant ribonuclease; bullfrog;
KW	signal peptide; cytotoxic fusion protein; cancer; autoimmune disease;
RNase.	
KW	
XX	
OS	Rana catesbeiana.
XX	
OS	Synthetic.
XX	
PN	WO9950398-A2.
XX	
PD	07-OCT-1999.
XX	
PF	26-MAR-1999; 99WO-US06641.
XX	
PR	27-MAR-1998; 98US-0079751.
XX	
PA	(USSH ) US DEPT HEALTH & HUMAN SERVICES.
XX	
PI	Newton DL, Rybak SM;
XX	
XX	WPI; 1999-610847/52.
DR	N-PSDB; AA208130.

XX New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases  
XX  
PS Claim 22; Page 62; 71pp; English.  
XX  
CC The present sequence is a Rana catesbeiana oocyte ribonuclease (RacOR1)  
CC protein encoded by a cDNA modified for expression in E. coli. Carboxy  
CC terminal end of RacOR1 has a covalently bound ligand binding moiety,  
CC which can be a IL2 antibody directed against CD22 on cancerous B cells  
CC or human chorionic gonadotropin (hCG) effective against Kaposi's  
CC Sarcoma cells. Recombinant ribonucleases can be expressed in bacteria  
CC without an N-terminal methionine due to the presence of a signal peptide  
CC that is cleaved by bacteria. The soluble expression of ribonucleases  
CC allows the proteins to be fused in-frame with ligand binding moieties to  
CC form cytotoxic fusion proteins. They can be used for treatment of cancer  
CC and autoimmune diseases.  
XX  
SQ Sequence 110 AA:  
Query Match 100.0%; Score 602; DB 20; Length 110;  
Best Local Similarity 100.0%; Pred. No. 2.3e-61;  
Matches 110; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 QNMTFQCKHINPIICNTIMDNIIYVGGCKRVNFTFISSATYKAICTGVINNVL 60  
DB 1 QNMTFQCKHINPIICNTIMDNIIYVGGCKRVNFTFISSATYKAICTGVINNVL 60  
QY 61 STTRFQNTCTRTSITPRCPYSSRTETNYICVGCENQYPVHFAGIGRCP 110  
DB 61 STTRFQNTCTRTSITPRCPYSSRTETNYICVGCENQYPVHFAGIGRCP 110  
RESULT 2  
ID AAY28873 standard; Protein: 111 AA.  
AC AAY28873:  
XX  
XX 25-JAN-2000 (first entry)  
DT  
XX  
DE Recombinant Met(-1) RacOR1.  
XX  
KW Recombinant Met(-1) Rana catesbeiana oocyte ribonuclease; RacOR1; CD22;  
KW covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell;  
KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;  
KW recombinant ribonuclease; cytotoxic fusion protein; cancer; bullfrog;  
KW Rnase; autoimmune disease.  
XX  
OS Rana catesbeiana.  
OS Synthetic.  
XX  
XX Key Location/Qualifiers  
FT Misc-difference 1 /note= "Met not found in wild type RacOR1"  
FT  
XX  
XX WO9950398-A2.  
XX  
XX 07-OCT-1999.  
XX  
XX 26-MAR-1999; 99WO-US06641.  
XX  
XX 27-MAR-1998; 98US-0079751.  
XX  
XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
XX  
XX Newton DL, Rybak SM;  
XX  
XX WPI: 1999-610847/52.  
XX  
XX N-PSDB; AA208131.  
XX  
XX New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases

XX Claim 22; Page 63; 71pp; English.  
PS  
XX  
CC The present sequence is a recombinant Rana catesbeiana oocyte  
CC ribonuclease (RacOR1) protein with Met at position 1. Carboxy terminal  
CC end of recombinant RacOR1 has a covalently bound ligand binding moiety,  
CC which can be a IL2 antibody directed against CD22 on cancerous B cells or  
CC human chorionic gonadotropin (hCG) effective against Kaposi's sarcoma  
CC cells. Recombinant ribonucleases can be expressed in bacteria without an  
CC N-terminal methionine due to the presence of a signal peptide that is  
CC cleaved by bacteria. The soluble expression of ribonuclease allows the  
CC proteins to be fused in-frame with ligand binding moieties to form  
CC cytotoxic fusion proteins. They can be used for treatment of cancer and  
CC autoimmune diseases.  
XX  
SQ Sequence 111 AA:  
Query Match 100.0%; Score 602; DB 20; Length 111;  
Best Local Similarity 100.0%; Pred. No. 2.3e-61;  
Matches 110; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 QNMTFQCKHINPIICNTIMDNIIYVGGCKRVNFTFISSATYKAICTGVINNVL 60  
DB 2 QNMTFQCKHINPIICNTIMDNIIYVGGCKRVNFTFISSATYKAICTGVINNVL 61  
QY 61 STTRFQNTCTRTSITPRCPYSSRTETNYICVGCENQYPVHFAGIGRCP 110  
DB 62 STTRFQNTCTRTSITPRCPYSSRTETNYICVGCENQYPVHFAGIGRCP 111  
RESULT 3  
ID AAY28877 standard; Protein: 110 AA.  
AC AAY28877:  
XX  
XX 25-JAN-2000 (first entry)  
DT  
XX  
DE Recombinant RacOR1 Gln1ser amino acid sequence.  
XX  
KW Recombinant Rana catesbeiana oocyte ribonuclease; RacOR1 Gln1ser; CD22;  
KW covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell;  
KW bullfrog; Kaposi's sarcoma; human chorionic gonadotropin; hCG; Rnase;  
KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;  
KW cancer; autoimmune disease.  
XX  
OS Rana catesbeiana.  
OS Synthetic.  
XX  
XX Key Location/Qualifiers  
FT Misc-difference 1 /note= "Wild type Gln replaced with Ser"  
FT  
XX  
XX WO9950398-A2.  
XX  
XX 07-OCT-1999.  
XX  
XX 26-MAR-1999; 99WO-US06641.  
XX  
XX 27-MAR-1998; 98US-0079751.  
XX  
XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
XX  
XX Newton DL, Rybak SM;  
XX  
XX WPI: 1999-610847/52.  
XX  
XX N-PSDB; AA208134.  
XX  
XX New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases  
PS Claim 22; Page 67; 71pp; English.

CC The present sequence is a recombinant Rana catesbeiana oocyte  
CC ribonuclease (RacOR1) protein with glutser. Carboxy terminal end of  
CC recombinant RacOR1 has a covalently bound ligand binding moiety, which  
CC can be a LL2 antibody directed against CD22 on cancerous B cells or  
CC human chorionic gonadotropin (hCG) effective against Kaposi's sarcoma  
CC cells. Recombinant ribonucleases can be expressed in bacteria without an  
CC N-terminal methionine due to the presence of a signal peptide that is  
CC cleaved by bacteria. The soluble expression of ribonuclease allows the  
CC proteins to be fused in-frame with ligand binding moieties to form  
CC cytotoxic fusion proteins. They can be used for treatment of cancer and  
CC autoimmune diseases.

XX Sequence 110 AA:

Query Match 99.2%; Score 597; DB 20; Length 110;  
Best Local Similarity 100.0%; Pred. No. 8.7e-61;  
Matches 109; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 NMATFOCKHILNPIICNTIMDNVYIVGQCKRVNFIISATVKAICTGVINMVL 61  
DB 2 NMATFOCKHILNPIICNTIMDNVYIVGQCKRVNFIISATVKAICTGVINMVL 61  
QY 62 TTRFQALNTCTRTSITPRCPYSSRTETNYICVKCENQYPVHFAIGRCP 110  
DB 62 TTRFQALNTCTRTSITPRCPYSSRTETNYICVKCENQYPVHFAIGRCP 110

## RESULT 4

AAI28878 ID AAY28878 standard; Protein: 111 AA.

XX AAY28878;

DT 25-JAN-2000 (first entry)

XX Recombinant Met(-1) RacOR1 glutser amino acid sequence.

XX Recombinant Met(-1) Rana catesbeiana oocyte ribonuclease glutser; RacOR1;  
KW covalently bound; LL2 antibody; ligand binding moiety; cancerous B cell;  
KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;  
KW recombinant ribonuclease; cytotoxic fusion protein; cancer; bullfrog;  
KW CD22; RNase; autoimmune disease.

XX Rana catesbeiana.  
OS Synthetic.

FH Key Location/Qualifiers

FT Misc-difference 1 /note= "Met not found in wild type RacOR1"

FT Misc-difference 2 /note= "Wild type Gln replaced with Ser"

PN W09950398-A2.

XX 07-OCT-1999.

PE 26-MAR-1999; 99WO-US06641.

PR 27-MAR-1998; 98US-0079751.

PA (USSH ) US DEPT HEALTH &amp; HUMAN SERVICES.

PI Newton DL, Rybak SM;

DR WPI: 1999-610847/52.

DR N-PSDB: AA208135.

PT New recombinant ribonucleases, used for killing target cells, e.g. for

PS treating cancers, viral infections or autoimmune diseases

XX Claim 22; Page 68; 71pp; English.

CC The present sequence is a recombinant Rana catesbeiana ribonuclease

CC (RacOR1) protein with Met at position 1 and glutser. Carboxy terminal end  
CC of recombinant RacOR1 has a covalently bound ligand binding moiety, which  
CC can be a LL2 antibody directed against CD22 on cancerous B cells or human  
CC chorionic gonadotropin (hCG) effective against Kaposi's sarcoma cells.  
CC Recombinant ribonucleases can be expressed in bacteria without an N-  
CC terminal methionine due to the presence of a signal peptide that is  
CC cleaved by bacteria. The soluble expression of ribonuclease allows the  
CC proteins to be fused in-frame with ligand binding moieties to form  
CC cytotoxic fusion proteins. They can be used for treatment of cancer and  
CC autoimmune diseases.

XX Sequence 111 AA:

Query Match 99.2%; Score 597; DB 20; Length 111;  
Best Local Similarity 100.0%; Pred. No. 8.8e-61;  
Matches 109; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 NMATFOCKHILNPIICNTIMDNVYIVGQCKRVNFIISATVKAICTGVINMVL 61  
DB 3 NMATFOCKHILNPIICNTIMDNVYIVGQCKRVNFIISATVKAICTGVINMVL 62  
QY 62 TTRFQALNTCTRTSITPRCPYSSRTETNYICVKCENQYPVHFAIGRCP 110  
DB 63 TTRFQALNTCTRTSITPRCPYSSRTETNYICVKCENQYPVHFAIGRCP 111

## RESULT 5

AAI28878 ID AAY28878 standard; Protein: 110 AA.

XX AAY28878;

DT 25-JAN-2000 (first entry)

XX Recombinant RacOR1 Met22Leu Met57Leu amino acid sequence.

XX Recombinant Rana catesbeiana oocyte ribonuclease; covalently bound;  
KW RacOR1 Met22Leu Met57Leu; LL2 antibody; ligand binding moiety; CD22;  
KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;  
KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;  
KW cancer; bullfrog; RNase; autoimmune disease.

XX Rana catesbeiana.  
OS Synthetic.

FH Key Location/Qualifiers

FT Misc-difference 22 /note= "Wild type Met replaced with Leu"

FT Misc-difference 57 /note= "Wild type Met replaced with Leu"

PN W09950398-A2.

XX 07-OCT-1999.

PE 26-MAR-1999; 99WO-US06641.

PR 27-MAR-1998; 98US-0079751.

PA (USSH ) US DEPT HEALTH &amp; HUMAN SERVICES.

PI Newton DL, Rybak SM;

DR WPI: 1999-610847/52.

DR N-PSDB: AA208132.

PT New recombinant ribonucleases, used for killing target cells, e.g. for

PS treating cancers, viral infections or autoimmune diseases

XX Claim 22; Page 64; 71pp; English.

CC The present sequence is a recombinant Rana catesbeiana oocyte

CC ribonuclease (RacOR1) protein with Met22Leu Met57Leu. Carboxy terminal

CC end of recombinant RacOR1 has a covalently bound ligand binding moiety,  
 CC which can be a LL2 antibody directed against CD22 on cancerous B cells,  
 CC or human chorionic gonadotrophin (hCG) effective against Kaposi's sarcoma  
 CC cells. Recombinant ribonucleases can be expressed in bacteria without an  
 CC N-terminal methionine due to the presence of a signal peptide that is  
 CC cleaved by bacteria. The soluble expression of ribonuclease allows the  
 CC proteins to be fused in-frame with ligand binding moieties to form  
 CC cytotoxic fusion proteins. They can be used for treatment of cancer and  
 CC autoimmune diseases.

XX Sequence 110 AA:  
 SQ

Query Match 99.0%; Score 596; DB 20; Length 110;  
 Best Local Similarity 98.2%; Pred. No. 1.1e-60;  
 Matches 108; Conservative 2; Mismatches 0; Indels 0; Gaps 0;

QY 1 QNMTFQOKHIINTPLICNTIMDNNTIYIGGCKRVNTFISSATVKAICTGVINLV 60  
 DB 1 QNMTFQOKHIINTPLICNTIMDNNTIYIGGCKRVNTFISSATVKAICTGVINLV 60

QY 61 STTRFQNLNCTRTSITPRPCPYSSRTETNYICVKCENQVPHFAGIGRCP 110  
 DB 61 STTRFQNLNCTRTSITPRPCPYSSRTETNYICVKCENQVPHFAGIGRCP 110

RESULT 6  
 AAY28876  
 ID AAY28876 standard; Protein: 111 AA.  
 XX AAY28876;  
 AC  
 XX  
 DT 25-JAN-2000 (first entry)  
 XX  
 DE Recombinant Met(-1) RacOR1 Met22Leu Met57Leu-(His)6 protein.  
 XX  
 XX Met(-1) Rana catesbeiana ribonuclease Met22Leu Met57Leu-(His)6; RacOR1;  
 KW recombinant; CD22; covalently bound; LL2 antibody; ligand binding moiety;  
 KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotrophin; hCG;  
 KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;  
 KW cancer; bullfrog; RNase; autoimmune disease.  
 XX  
 XX Rana catesbeiana.  
 OS Synthetic.  
 OS  
 XX  
 FH Key Location/Qualifiers  
 FT MISC-difference 1 /note= "(His)6 histidine tag attached to N-terminal Met"  
 FT MISC-difference 1 /note= "Met not found in wild type RacOR1"  
 FT MISC-difference 23 /note= "Wild type Met replaced with Leu"  
 FT MISC-difference 58 /note= "Wild type Met replaced with Leu"  
 FT  
 XX  
 XX W09950398-A2.  
 PN  
 XX  
 XX 07-OCT-1999.  
 PD  
 XX  
 XX 26-MAR-1999; 99WO-US06641.  
 PF  
 XX  
 XX 27-MAR-1998; 98US-0079751.  
 PR  
 XX  
 XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PA  
 XX  
 XX Newton DL, Rybak SM;  
 PI  
 XX  
 XX WPI: 1999-610847/52.  
 DR  
 XX  
 XX N-PSDB; AA208133.  
 DR  
 XX  
 XX New recombinant ribonucleases, used for killing target cells, e.g. for  
 PT treating tumors, viral infections or autoimmune diseases  
 PT  
 XX  
 PS Claim 22; Page 66; 71pp; English.

XX  
 CC The present sequence is a recombinant Rana catesbeiana oocyte  
 CC ribonuclease (RacOR1) protein with Met at position 1 attached to a  
 CC (His)6 tag, Met23Leu and Met58Leu. Carboxy terminal end of recombinant  
 CC RacOR1 has a covalently bound ligand binding moiety, which can be a LL2  
 CC antibody directed against CD22 on cancerous B cells or human chorionic  
 CC gonadotrophin (hCG) effective against Kaposi's sarcoma cells. Recombinant  
 CC ribonucleases can be expressed in bacteria without an N-terminal  
 CC methionine due to the presence of a signal peptide that is cleaved by  
 CC bacteria. The soluble expression of ribonuclease allows the proteins to  
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
 CC proteins. They can be used for treatment of cancer and autoimmune  
 CC diseases.

XX Sequence 111 AA:  
 SQ

Query Match 99.0%; Score 596; DB 20; Length 111;  
 Best Local Similarity 98.2%; Pred. No. 1.1e-60;  
 Matches 108; Conservative 2; Mismatches 0; Indels 0; Gaps 0;

QY 1 QNMTFQOKHIINTPLICNTIMDNNTIYIGGCKRVNTFISSATVKAICTGVINLV 60  
 DB 2 QNMTFQOKHIINTPLICNTIMDNNTIYIGGCKRVNTFISSATVKAICTGVINLV 61

QY 61 STTRFQNLNCTRTSITPRPCPYSSRTETNYICVKCENQVPHFAGIGRCP 110  
 DB 62 STTRFQNLNCTRTSITPRPCPYSSRTETNYICVKCENQVPHFAGIGRCP 111

RESULT 7  
 AAY33321  
 ID AAY33321 standard; Protein: 111 AA.  
 XX AAY33321;  
 AC  
 XX  
 DT 29-NOV-1999 (first entry)  
 XX  
 DE Frog lectin protein fragment.  
 XX  
 XX Cytotoxic; RNase; ribonuclease; pancreatic; antibody; light chain;  
 KW heavy chain; cell surface marker; treatment; tumor; viral infection;  
 KW parasite infection; immune dysfunctional cell; autoimmune disease;  
 KW contraceptive; cell separation; transplantation; bone marrow ablation;  
 KW leukemia cell; T-cell; graft-versus-host disease; bullfrog; lectin.  
 XX  
 XX Rana catesbeiana.  
 OS  
 OS  
 XX  
 PN US9595073-A.  
 PN  
 XX  
 PD 21-SEP-1999.  
 PD  
 XX  
 XX 09-JUL-1997; 97US-0891848.  
 PF  
 XX  
 XX 22-SEP-1993; 93US-0125462.  
 PR  
 XX 22-OCT-1991; 91US-0779195.  
 PR 20-APR-1990; 90US-0510696.  
 PR 04-FEB-1993; 93US-0014082.  
 PR  
 XX  
 XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PA  
 XX  
 XX Rybak SM, Newton DL, Nicholls PJ, Youle RJ;  
 PI  
 XX  
 XX WPI: 1999-560488/47.  
 DR  
 XX  
 XX Recombinantly fused pancreatic RNase-targeting proteins useful for  
 PT treating tumors, infections, immune or autoimmune disorders and as a  
 PT contraceptive  
 PT  
 XX  
 XX Example 3; Fig 19; 47pp; English.  
 PS  
 XX  
 XX This invention describes a novel nucleic acid construct comprising  
 CC sequences encoding functional pancreatic RNase and a second protein  
 CC (preferably the light and heavy chains of an antibody) which binds a

CC specific cell surface marker on a target cell and functions as a  
 CC cytotoxic agent. The products can be used for selectively killing cells  
 CC expressing a specific surface marker. They can be used for treating  
 CC tumors or infected cells (e.g. cells infected by viruses (especially  
 CC latent or chronic virus infections, such as human immunodeficiency virus  
 CC (HIV-1, Epstein-Barr virus, herpes viruses (herpes simplex types 1 and  
 CC 11), hepatitis viruses (B, non-A-non-B, and delta), herpes zoster,  
 CC cytomegalovirus) and cells infected with parasites (such as the malaria  
 CC parasite). They can also be used for treating immune dysfunctional cells  
 CC in immune and autoimmune diseases. Additionally, they may be used as  
 CC contraceptives. Finally they can also be used for cell separation in  
 CC vitro by selectively killing unwanted types of cells (e.g. in bone  
 CC marrow) prior to transplantation into a patient undergoing marrow  
 CC ablation by radiation or for killing leukemia cells or T-cells that would  
 CC cause graft-versus-host disease. This sequence represents a fullfrog  
 CC (Rana catesbeiana) lectin used to describe the method of the invention.

XX Sequence 111 AA:

Query Match 97.8%; Score 588.5; DB 20; Length 111;

Best Local Similarity 98.2%; Pred. No. 8.3e-60;

Matches 109; Conservative 1; Mismatches 0; Indels 1; Gaps 1;

OY 1 QNNAFPOQKHIIINTPII-CNTIMDNIIYVGGCKRNVFTISSATTVAICGVI-NMN 59

DB 1 ENNAFPOQKHIIINTPII-CNTIMDNIIYVGGCKRNVFTISSATTVAICGVI-NMN 60

OY 60 LSTTRPOLNCTRTSTPRPCPSRTEFTYICVGCENOPVHPAGIGRC 110

DB 61 LSTTRPOLNCTRTSTPRPCPSRTEFTYICVGCENOPVHPAGIGRC 111

RESULT 8

AAW06544 standard; protein: 104 AA.

AC AAW06544:

DT 22-AUG-1997 (first entry)

DE Antitumour protein from Rana pipiens oocytes.

KW Tumour; chemotherapy; radiotherapy; frog.

OS Rana pipiens.

PN W09639428-A1.

PD 12-DEC-1996.

PF 03-JUN-1996; 96W0-US08304.

PR 06-JUN-1995; 95US-0467955.

PA (ALFA-) ALFACELL CORP.

PI Ardelet WJ;

DR WPI; 1997-043063/G4.

PT Antitumour proteins from Rana pipiens oocyte(s) - have fewer

PT disadvantages than chemotherapy, surgery and radiotherapy

PS Claim 8; Page 28; 45pp: English.

CC The present sequence is a specifically claimed example of an  
 CC antitumour protein from the generic protein in AAW18224, with the  
 CC molecular weight 12000. This is one of two preferred proteins (the  
 CC other in AAW06543) that have been isolated from Rana pipiens oocytes.  
 CC Both proteins have a blocked amino terminal group and are essentially  
 CC free of carbohydrates. The proteins are used to treat tumours. Use of  
 CC the peptides has fewer disadvantages than chemotherapy, radiotherapy  
 CC and surgery in the treatment of tumours.

XX Sequence 104 AA:

Query Match 47.3%; Score 284.5; DB 18; Length 104;

Best Local Similarity 49.5%; Pred. No. 6.2e-25;

Matches 55; Conservative 16; Mismatches 31; Indels 9; Gaps 4;

OY 1 QNNAFPOQKHIIINTPII-CNTIMDNIIYVGGCKRNVFTISSATTVAICGVI-NMN 58

DB 1 EDLTFQKKHVTNRVDNCNINMSTNLF---HCKDKNFTIYSRPPVKAICGIIASKN 56

OY 59 VLTSTRPOLNCTRTSTPRPCPSRTEFTYICVGCENOPVHPAGIGRC 109

DB 57 VLTSTRPOLNCTRTSTPRPCPSRTEFTYICVGCENOPVHPAGIGRC 104

RESULT 9

AAV28865 standard; protein: 104 AA.

AC AAV28865:

DT 25-JAN-2000 (first entry)

DE Rana pipiens liver ribonuclease (RaplR1).

KW Rana pipiens liver ribonuclease; RapLR1; covalently bound; LL2 antibody;

KW ligand binding moiety; CD22; cancerous B cell; Kaposi's Sarcoma; frog;

KW human chorionic gonadotropin; hCG; recombinant ribonuclease; RNase;

KW signal peptide; cytotoxic fusion protein; cancer; autoimmune disease.

OS Rana pipiens.

PN W09950398-A2.

PD 07-OCT-1999.

PF 26-MAR-1999; 99W0-US06641.

PR 27-MAR-1998; 98US-0079751.

PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.

PI Newton DL, Rybak SM;

DR WPI; 1999-610847/52.

DR N-PSDB; AAZ08124.

PT New recombinant ribonucleases, used for killing target cells, e.g. for

PT treating cancers, viral infections or autoimmune diseases

PS Claim 1; Page 55; 71pp: English.

CC The present sequence is Rana pipiens liver ribonuclease (RaplR1)  
 CC protein. Carboxy terminal end of RapLR1 has a covalently bound  
 CC ligand binding moiety, which can be a LL2 antibody directed against  
 CC CD22 on cancerous B cells or human chorionic gonadotropin (hCG)  
 CC effective against Kaposi's Sarcoma cells. Recombinant ribonucleases can  
 CC be expressed in bacteria without an N-terminal methionine due to the  
 CC presence of a signal peptide that is cleaved by bacteria. The soluble  
 CC expression of ribonuclease allows the proteins to be fused in-frame with  
 CC ligand binding moieties to form cytotoxic fusion proteins. They can be  
 CC used for treatment of cancer and autoimmune diseases.

XX Sequence 104 AA:

Query Match 46.8%; Score 281.5; DB 20; Length 104;

Best Local Similarity 49.5%; Pred. No. 1.4e-24;

Matches 55; Conservative 15; Mismatches 32; Indels 9; Gaps 4;

OY 1 QNNAFPOQKHIIINTPII-CNTIMDNIIYVGGCKRNVFTISSATTVAICGVI-NMN 58

DB 1 QDMLTFQKKHIIINTPII-CNTIMDNIIYVGGCKRNVFTISSATTVAICGVI-NMN 56

OY 59 VLSTTRQLNTCTRTSTTPRCPPYSSRTETNYICVKGCEQYVPHFAGIGRC 109  
 DB 57 VLTTSERFLSDC---NWTSRPCKYKLLKSTNTFCVTCENQAPVHFVGCHC 104

## RESULT 10

AAy28867  
 ID AAY28867 standard; Protein; 105 AA.

AC AAY28867;

DT 25-JAN-2000 (first entry)

DE Recombinant Met(-1) RAPLRI.

KW Recombinant Met(-1) Rana pipiens ribonuclease; RAPLRI; CD22; RNase;  
 KW covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell;  
 KW Kaposi's sarcoma; human chorionic gonadotrophin; hCG; signal peptide;  
 KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;  
 KW autoimmune disease.

OS Rana pipiens.  
 OS Synthetic.

FT Key Location/Qualifiers

FT Misc-difference 1 /note="Met not found in wild type RAPLRI"

PN W09950398-AZ.

PD 07-OCT-1999.

PF 26-MAR-1999; 99WO-US06641.

PR 27-MAR-1998; 98US-0079751.

PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.

PI Newton DL, Rybak SM;

DR WPI; 1999-610847/52.

DR N-PSDB; AA08136.

PT New recombinant ribonucleases, used for killing target cells, e.g. for  
 PT treating cancers, viral infections or autoimmune diseases

PS Claim 34; Page 57; 71pp; English.

CC The present sequence is a recombinant Rana pipiens ribonuclease (RAPLRI)  
 CC protein with Met at position 1. Carboxy terminal end of recombinant  
 CC RAPLRI has a covalently bound ligand binding moiety, which can be a IL2  
 CC gonadotrophin (hCG) effective against CD22 on cancerous B cells or human chorionic  
 CC ribonucleases can be expressed in bacteria without an N-terminal  
 CC methionine due to the presence of a signal peptide that is cleaved by  
 CC bacteria. The soluble expression of ribonuclease allows the proteins to  
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
 CC proteins. They can be used for treatment of cancer and autoimmune  
 CC diseases.

SO Sequence 105 AA;

Query Match 46.8%; Score 281.5; DB 20; Length 105;  
 Best Local Similarity 49.5%; Pred. No. 1.4e-24;

Matches 55; Conservative 15; Mismatches 32; Indels 9; Gaps 4;

OY 1 QNATFOQKHINT-PIICNTIMDNNTIYVGGQCKRVNFTISSATVKAICGVI-NMN 58  
 DB 2 QDWLTFOQKHINTRDVDCNNIMSTNLF---HCKDKNTFIYSRPEPVKAICGIISK 57  
 OY 59 VLSTTRQLNTCTRTSTTPRCPPYSSRTETNYICVKGCEQYVPHFAGIGRC 109

DB 58 VLTTSERFLSDC---NWTSRPCKYKLLKSTNTFCVTCENQAPVHFVGCHC 105

## RESULT 11

AAy28879  
 ID AAY28879 standard; Protein; 127 AA.

AC AAY28879;

DT 25-JAN-2000 (first entry)

DE Rana pipiens Clone 5a1b ribonuclease.

KW Rana pipiens ribonuclease Clone 5a1b; RAPLRI; covalently bound; RNase;  
 KW IL2 antibody; ligand binding moiety; CD22; cancerous B cell; onconase;  
 KW Kaposi's Sarcoma; human chorionic gonadotrophin; hCG; cancer;  
 KW recombinant ribonuclease; frog; signal peptide; cytotoxic fusion protein;  
 KW autoimmune disease.

OS Rana pipiens.

FT Key Location/Qualifiers

FT Peptide 1..23 /label=Signal-peptide

FT Protein 24..127 /note="Putative" /label= Rana\_pipiens\_Clone\_5a1b\_ribonuclease

PN W09950398-AZ.

PD 07-OCT-1999.

PF 26-MAR-1999; 99WO-US06641.

PR 27-MAR-1998; 98US-0079751.

PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.

PI Newton DL, Rybak SM;

DR WPI; 1999-610847/52.

DR N-PSDB; AA08136.

PT New recombinant ribonucleases, used for killing target cells, e.g. for  
 PT treating cancers, viral infections or autoimmune diseases

PS Disclosure; Page 69; 71pp; English.

CC The present sequence is a Rana pipiens Clone 5a1b ribonuclease (RAPLRI).  
 CC It is encoded by Clone 5a1b cDNA obtained from Rana pipiens liver mRNA  
 CC library. It exhibits differences with Onconase (RPM) at amino acid  
 CC residues 11, 20, 85 and 103. Carboxy terminal end of RAPLRI has a  
 CC covalently bound ligand binding moiety, which can be a IL2 antibody  
 CC directed against CD22 on cancerous B cells or human chorionic  
 CC gonadotrophin (hCG) effective against Kaposi's Sarcoma cells. Recombinant  
 CC ribonucleases can be expressed in bacteria without an N-terminal  
 CC methionine due to the presence of a signal peptide that is cleaved by  
 CC bacteria. The soluble expression of ribonuclease allows the proteins to  
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
 CC proteins. They can be used for treatment of cancer and autoimmune  
 CC diseases.

SO Sequence 127 AA;

Query Match 46.8%; Score 281.5; DB 20; Length 127;  
 Best Local Similarity 49.5%; Pred. No. 1.8e-24;

Matches 55; Conservative 15; Mismatches 32; Indels 9; Gaps 4;

OY 1 QNATFOQKHINT-PIICNTIMDNNTIYVGGQCKRVNFTISSATVKAICGVI-NMN 58  
 DB 24 QDWLTFOQKHINTRDVDCNNIMSTNLF---HCKDKNTFIYSRPEPVKAICGIISK 79  
 OY 59 VLSTTRQLNTCTRTSTTPRCPPYSSRTETNYICVKGCEQYVPHFAGIGRC 109

Db 80 VLTSEFLSDC---NVTSRPCKYKLLKSTNTEFCVTCENQAPVHVGVC 127

## RESULT 12

AAV28866 ID AAV28866 standard; Protein: 104 AA.

AC AAV28866; XX

DT 25-JAN-2000 (first entry)

XX Recombinant RapLRI Met23leu amino acid sequence.

XX Recombinant Rana pipiens ribonuclease; RapLRI Met23leu; covalently bound;

KW IL2 antibody; ligand binding moiety; CD22; cancerous B cell; RNase;

KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;

KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;

XX autoimmune disease.

XX Rana pipiens.

OS Synthetic.

XX Key

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

FT Misc-difference 23

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FT Misc-difference 23

RESULT 13  
AAV28869 ID AAV28869 standard; Protein: 105 AA.

AC AAV28869; XX

DT 25-JAN-2000 (first entry)

XX Recombinant Met(-1) RapLRI Met23leu-(His)6 protein.

XX Recombinant Met(-1) Rana pipiens ribonuclease Met23leu-(His)6; RapLRI;

KW CD22; covalently bound; IL2 antibody; ligand binding moiety; RNase;

KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;

KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;

KW cancer; frog; autoimmune disease.

XX Rana pipiens.

OS Synthetic.

XX Key

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

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Query Match 46.3%; Score 278.5; DB 20; Length 105;  
Best Local Similarity 48.6%; Pred. No. 31e-24;  
Matches 54; Conservative 16; Mismatches 32; Indels 9; Gaps 4;

AC AAV28869; XX

DT 25-JAN-2000 (first entry)

XX Recombinant Met(-1) RapLRI Met23leu-(His)6 protein.

XX Recombinant Met(-1) Rana pipiens ribonuclease Met23leu-(His)6; RapLRI;

KW CD22; covalently bound; IL2 antibody; ligand binding moiety; RNase;

KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;

KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;

KW cancer; frog; autoimmune disease.

XX Rana pipiens.

OS Synthetic.

XX Key

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

FT Misc-difference 1

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FT Misc-difference 1

FT Misc-difference 1

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FT Misc-difference 1

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XX      Key          Location/Qualifiers
FH      Modified-size    1
FT      /note="this Gln is autocyclised to pyroglutamic acid"
XX      US6175003-BI.
XX      PD            16-JAN-2001.
XX      PF            10-SEP-1999;   99US-0394268.
XX      PR            10-SEP-1999;   99US-0394268.
PA      (ALFA-) ALFACELL CORP.
XX      Saxena SK;
XX      WPI; 2001-167808/17.
PT      New nucleic acids encoding a ribonuclease (Rnase), useful for the
XX      precise targeting of Rnase to a predetermined cell receptor .
PS      Claim 1; Columns 5-6; 7pp; English.

The present sequence represents a frog ribonuclease protein (ranpinase)
(Rnase). The specification describes a synthetic ribonuclease protein,
in which the addition of cysteine in the ribonuclease facilitates the
chemical linking of a targeting molecule by the single reactive
sulfhydryl group. The specification also describes a method for the
production of ranpinase using DNA technology instead of processing
biological material. The re-engineering of the protein molecule allows
easier attachment to a targeting molecule thereby making it possible for
the ribonuclease to be delivered to a particular cell receptor where it
might be most effective.

SQ      Sequence      104 AA:

Query Match           46.1%; Score 277.5; DB 22; Length 104;
Best Local Similarity 49.5%; Pred No. 4e-24;
Matches 55; Conservative 15; Mismatches 32; Indels 9; Gaps 4.

OY      1 QNMAFDQKHIIINT-PIICNTIMDNNIIYVGQCKRVMTFIISATVKAICTGVI-MWN 58
       1 :| | | | | | | | : | : | | | : | | | | | | | | | | | | | |
Db      1 QDWLTFQKHHITNRDVDCDNIMSTNFL----HCKDKMTFIYSRREPYPVKAICKGITASKN 56
       1 :| | | | | | | | : | : | | | : | | | | | | | | | | | | | |

OY      59 VLSTTRQLNCTRTSITPRCPPISSRTETNYICVKCENOYPVAHFAGIGRC 109
       || | : | | : | : | | | | : | | | | | | | | | | | | | |
Db      57 VLTTSERYLSDC---NVTSRPCCKYKLKSTNKFCVTCENOAPVHFVGVGCSC 104
       || | : | | : | : | | | | : | | | | | | | | | | | | | |

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